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Amendments to the Claims

This listing of claims replaces all prior versions and listings of claims in the application.

Listing of the Claims:

1. (Currently amended) A purified [[ICOS]] polypeptide having altered affinity for B7-H2 compared to a wild-type ICOS polypeptide, consisting of:

(a) a variant of

- (i) a wild-type ICOS amino acid sequence consisting of an extracellular domain of wild-type ICOS, the wild-type ICOS extracellular domain being SEQ ID NO:10 or SEQ ID NO:9, or
- (ii) a wild-type ICOS amino acid sequence consisting of a fragment of at least 8 amino acids of the extracellular domain,

the variant:

consisting of an amino acid sequence that differs by one or more amino acid substitutions from, but is at least 75% homologous to, its corresponding wild-type ICOS amino acid sequence; and

having altered affinity for B7-H2 compared to its corresponding wild-type ICOS amino acid sequence, wherein said altered affinity for B7-H2 is at least 6% of the affinity of [[said]] the corresponding wild-type ICOS [[polypeptide]] amino acid sequence; or

(b) the variant of (a) and: (I) a peptide sequence unrelated to ICOS attached to the N-terminus of the variant of (a); (II) a peptide sequence unrelated to ICOS attached to the C-terminus of the variant of (a); or (III) a peptide sequence unrelated to ICOS attached to the N-terminus of the variant of (a) and a peptide sequence unrelated to ICOS attached to the C-terminus of the variant of (a).

2. (Cancelled)

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3. (Currently amended) The purified [[ICOS]] polypeptide of claim 2, wherein said difference is the variant differs from its corresponding wild-type amino acid sequence at [[amino acid]] a position corresponding to amino acid 76 of SEQ ID NO:12.

- 4. (Currently amended) The purified [[ICOS]] polypeptide of claim 3, wherein, in the variant, the amino acid at the position corresponding to said amino acid [[position]] 76 of SEQ ID NO:12 [[contains a]] is glutamine.
- 5. (Currently amended) The purified [[ICOS]] polypeptide of claim 2, wherein said difference is the variant differs from its corresponding wild-type amino acid sequence at [[amino acid]] a position corresponding to amino acid 52 of SEQ ID NO:12.
- 6. (Currently amended) The purified [[ICOS]] polypeptide of claim 5, wherein, in the variant, the amino acid at the position corresponding to said amino acid [[position]] 52 of SEQ ID NO:12 [[contains a]] is serine.
- 7. (Currently amended) The purified [[ICOS]] polypeptide of claim 1, wherein said [[polypeptide]] variant is capable of inhibiting T cell activation in a T cell proliferation assay.
- 8. (Withdrawn) An isolated nucleic acid molecule comprising a nucleic acid sequence that encodes the polypeptide of claim 1.
- 9. (11). (Cancelled)
- 12. (Currently amended) A method for inhibiting T cell activation, comprising contacting an antigen-presenting cell with [[a]] the purified [[ICOS]] polypeptide of claim 1, wherein said polypeptide is capable of binding to B7-H2 with increased affinity relative to [[a]] its corresponding wild-type ICOS [polypeptide having the amino acid sequence of SEQ ID NO:12.

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13. (Currently amended) The method of claim 12, wherein said purified ICOS polypeptide variant comprises a Ser76Glu mutation.

- 14. (Currently amended) The method of claim 12, wherein said purified ICOS polypeptide variant comprises a Lys52Ser mutation.
- 15. (Currently amended) A method for inhibiting T cell activation in a subject, comprising administering to the subject an amount of the purified [[ICOS]] polypeptide of claim 1 that is capable of inhibiting a T cell response in said subject.
- 16. (Currently amended) The method of claim 15, wherein said [[ICOS polypeptide]] <u>variant</u> comprises a Ser76Glu mutation.
- 17. (Currently amended) The method of claim 15, wherein said [[ICOS polypeptide]] variant comprises a Lys52Ser mutation.
- 18. (Withdrawn) The method of claim 15, wherein said subject has an autoimmune disease.
- 19. (Withdrawn) The method of claim 18, wherein said subject has rheumatoid arthritis.
- 20. (Withdrawn) The method of claim 18, wherein said subject has systemic lupus erythematosus.
- 21. (Withdrawn) The method of claim 18, wherein said subject has diabetes mellitus.
- 22. (Withdrawn) The method of claim 15, wherein said subject is a transplant recipient.
- 23. (Withdrawn) A method for making an ICOS polypeptide, comprising culturing the cell of claim 11 and isolating said ICOS polypeptide from said culture.
- 24. (New) The purified polypeptide of claim 1, wherein the peptide sequence unrelated to the ICOS or the second peptide sequence unrelated to ICOS is a blocking agent that facilitates survival of the polypeptide *in vivo*.

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25. (New) The purified polypeptide of claim 1, wherein the peptide sequence unrelated to the ICOS or the second peptide sequence unrelated to ICOS is a tag amino acid sequence.

- 26. (New) The purified polypeptide of claim 1, wherein the peptide sequence unrelated to the ICOS or the second peptide sequence unrelated to ICOS is an immunoglobulin Fc fragment sequence.
- 27. (New) The purified polypeptide of claim 1, wherein said altered affinity is increased affinity.